PATENT COOPERATION TREA.

PCT



INTERNATIONAL SEARCH REPORT

Indicants or agents file reference	(PCT Article 18 and Rules 43 and 44)	Bu
pplicant's or agent's file reference	FOR FURTHER see Notification of (Form PCT/ISA/2	of Transmittal of International Search Report 20) as well as, where applicable, item 5 below.
ternational application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)
CT/US 99/26772	09/11/1999	10/11/1998
plicant		
AXYGEN, INC. et al.		
This International Search Report has be	een prepared by this International Searching Auth transmitted to the International Bureau.	nority and is transmitted to the applicant
according to Article 16. A copy is being		
This International Search Report consi	sts of a total of sheets.	
It is also accompanied	by a copy of each prior art document cited in this	report.
. Basis of the report		
•	he international search was carried out on the bas	sis of the international application in the
	unless otherwise indicated under this item.	
the international search Authority (Rule 23.1(b)	h was carried out on the basis of a translation of the).	he international application furnished to this
	and/or amino acid sequence disclosed in the in	ternational application, the international search
was carried out on the basis of contained in the internation	the sequence listing : ational application in written form.	
	nternational application in computer readable form	n.
furnished subsequently	y to this Authority in written form.	
	y to this Authority in computer readble form.	
the statement that the	subsequently furnished written sequence listing d n as filed has been furnished.	oes not go beyond the disclosure in the
	information recorded in computer readable form is	s identical to the written sequence listing has bee
. X Certain claims were t	ound unsearchable (See Box I)	
Unity of Invention is		
. With regard to the title ,		
	s submitted by the applicant.	
	blished by this Authority to read as follows:	
	,5-BISPHOSPHATE CARBOXYLASE/O	XYGENASE
 With regard to the abstract, 		
X the text is approved as	s submitted by the applicant.	
the text has been esta within one month from	blished, according to Rule 38.2(b), by this Authori the date of mailing of this international search re	ity as it appears in Box III. The applicant may, port, submit comments to this Authority.
	oublished with the abstract is Figure No.	
as suggested by the a		None of the figures.
because the applicant	failed to suggest a figure.	

because this figure better characterizes the invention.

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 99/26772

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This Int	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
17 7	because they relate to subject matter not required to be searched by this Authority, namely: 15–18 (incomplete)
<u> </u>	because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically: see FURTHER INFORMATION sheet PCT/ISA/210
з. [Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
1.	
<u></u>	searchable claims. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee. As only some of the required additional search fees were timely paid by the applicant, this International Search Report



FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Claims Nos.: 15-18 (incomplete)

Present claims 15-18 relate to a polynucleotide comprising a gene characterised by the expression "shuffled", which describes a process of nucleic acid rearrangment that may either occur naturally during evolution (see Wolter F.P. et al., Plant Physiol., 1997, 114 (1), p.131-6) or may be carried out by man as claimed in present claims 1-9. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Consequently, the search has been carried out for those parts of the claims which appear to be clear and disclosed, namely those parts relating to a polynucleotide comprising a sequence encoding a Rubisco Form I L subunit gene shuffled by a method as claimed in claims 1-9 and linked to a selectable marker gene and further optionally linked to upstream and downstream flanking recombinogenic sequences, whereby said polynucleotide codes for an enzyme with a Km for CO2 (O2) which is significantly lower (higher) than the respective Km of any naturally-occurring rubisco enzyme.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

INTERNATIONAL SEARCH REPORT

iternational Application No PCT/US 99/26772

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12N9/88 C12N15/29

C12Q1/68

C12N15/82

C12N5/14

C12P19/34

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

C12N C12P C12Q

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

c data base consulted during the international search (name of data base and, where practical, search terms used)

JUL 1 7 2000

C. DOCUMENTS CONSIDERE	ED TO BE RELEVANT
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Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	SPREITZET R.J.: "GENETIC DISSECTION OF RUBISCO STRUCTURE AND FUNCTION" ANNUAL REVIEW OF PLANT PHYSIOLOGY AND PLANT MOLECULAR BIOLOGY, vol. 44, 1993, pages 411-434, XP000864484	1-4, 10-12, 15-18
X	page 412, line 15 - line 34 page 415, line 6 -page 423, line 42	15-18
Y	WO 97 35966 A (MAXYGEN INC.) 2 October 1997 (1997-10-02)	1-4, 10-12,
	page 47, line 19 - line 22; claims 1,5,12-19	15-21
	-/	

Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
"A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "&" document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
28 March 2000	12/04/2000
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer Schönwasser, D

IN TRNATIONAL SEARCH REPORT

International Application No

C.(Continu	ustion) DOCUMENTS CONSIDERED TO BE RELEVANT	PCT/US 99/26772
Category °		Relevant to claim No.
 Y		Treisvant to Ciaim No.
	LARIMER F.W. ET AL.: "Rhodobacter capsulatus Calvin cycle carbon dioxide fixation operon: fructose-1,6-/sedoheptulose-1,7-bisphospha te aldolase (cbbA) gene, partial cds, Form II ribulose-1,5-bisphosphate carboxylase/oxygenase (cbbM) gene, complete cds, and Calvin cycle operon: pentose-5-phosphate-3-epimerase (cbbE), etc." EMBL DATABASE ENTRY RC23145; ACCESSION NO. U23145,14 April 1995 (1995-04-14), XP002132955	19-21
	CRAMERI A ET AL: "DNA SHUFFLING OF A FAMILY OF GENES FROM DIVERSE SPECIES ACCELERATES DIRECTED EVOLUTION" NATURE, vol. 391, 15 January 1998 (1998-01-15), pages 288-291, XP000775869 ISSN: 0028-0836 page 288, column 2, line 24 - line 28 page 289, column 1, line 4 -page 290, column 1, line 2	1-26
	JAMET E. ET AL.: "Genes encoding the small subunit of RUBISCO belong to two highly conserved subfamilies in Nicotianeae" JOURNAL OF MOLECULAR EVOLUTION, vol. 33, no. 3, September 1991 (1991-09), pages 226-236, XP000864454 the whole document	1-26
	WOLTER F.P. ET AL.: "rbcS genes in Solanum tuberosum: Conservation of transit peptide and exon shuffling during evolution" PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF USA, vol. 85, February 1988 (1988-02), pages 846-850, XP000877412	1-26
	FLACHMANN R. ET AL.: "Mutations in the Small Subunit of Ribulose-1,5-Bisphosphate Carboxylase/Oxygenase Increase the Formation of the Misfire Product Xylulose-1,5-Bisphosphate" PLANT PHYSIOLOGY, vol. 114, no. 1, May 1997 (1997-05), pages 131-136, XP000877382	1-26

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information on patent family members

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			ΑU	2337797 A	17-10-1997
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			JP 20	00500981 T	02-02-2000
			WO	9720078 A	05-06-1997

